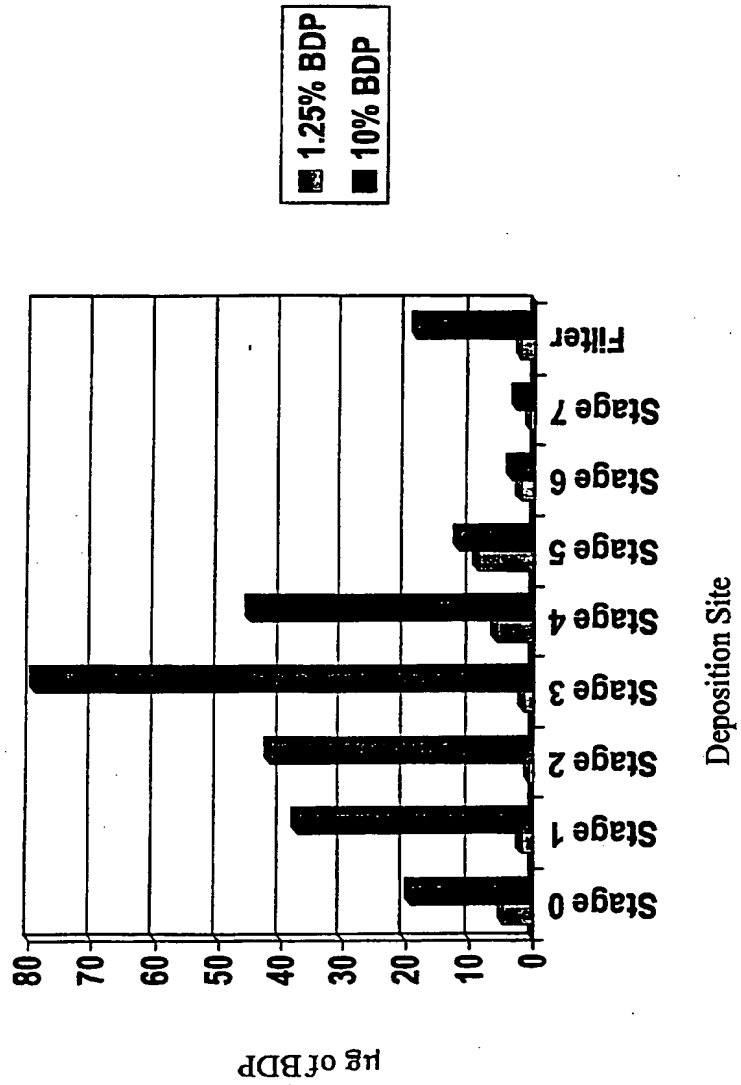


Title: LIQUID DROPLET AEROSOLS OF
NANOPARTICULATE DRUGS
Inventor(s): Bosch et al.
Appl. No.: 09/597,738

FIGURE 1
In Vitro Deposition Pattern of
Aerosolized BDP Dispersions



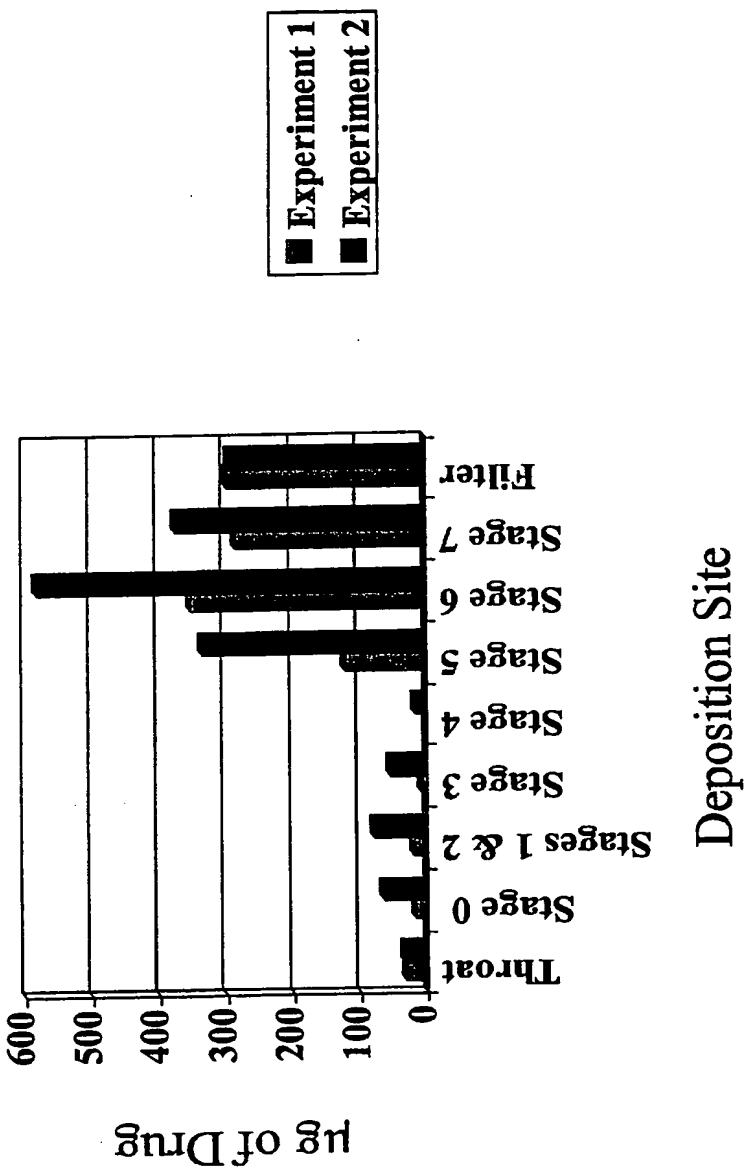
SEP 28 2004
PATENT & TRADEMARK CO.

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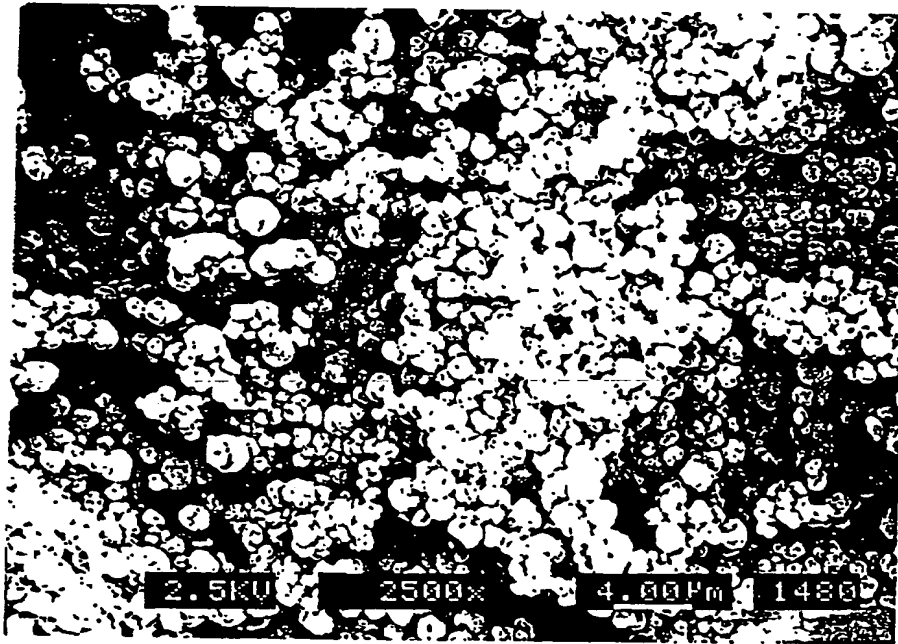
FIGURE 2
In Vitro Deposition Pattern of
Nanoparticulate BDP Suspensions





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FIGURE 4



Spray Dried Nanoparticulate Naproxen

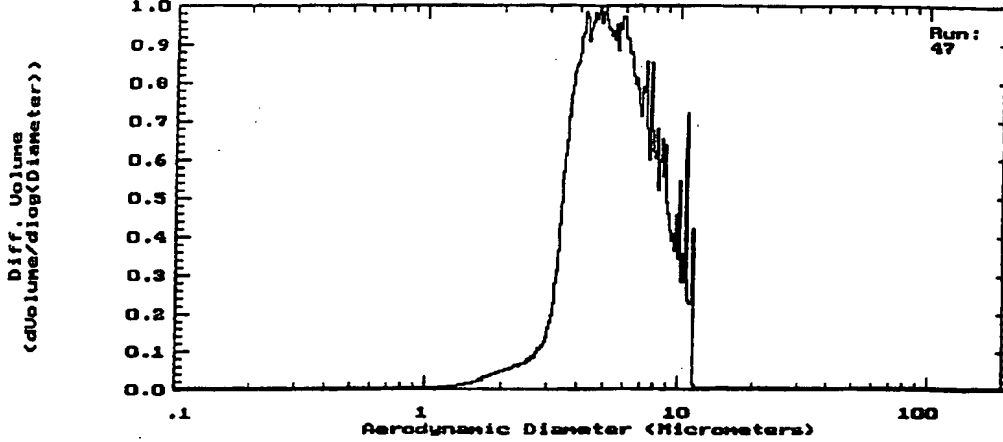
[illegible]



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FIGURE 6

API AEROSIZER-LD U7.10.09



spray dried ta

STATISTICS			PARAMETERS		UNDER	SIZE	UNDER	SIZE
Mean Size	: 5.540		Material	: SDI-naproxen	10%	3.600	90%	9.082
Standard Deviation	: 1.455		Density	: 1.26	50%	5.516		
D(4,3)	: 5.924		Run Length (sec)	: 189.6				
D(3,2)	: 5.146		PMT Voltage	: 1100.0				
Mode (Log Scale)	: 4.82		Sum of channels	: 100494				
Specific Surface Area	: 0.93	sq meter/g	Lower Size Limit	: 0.10				
			Upper Size Limit	: 200.00				
			Nozzle Type	: 700um				
			Baseline Offset	: 0.10				
			Noise Filter	: 6.00				
			Regularization	: Off				

UPPER SIZE	% IN	LOWER SIZE	% UNDER	UPPER SIZE	% IN	LOWER SIZE	% UNDER	UPPER SIZE	% IN	LOWER SIZE	% UNDER	UPPER SIZE	% IN	LOWER SIZE	% UNDER
		100	0.0000	86.0	100.00	10.0	7.5026	8.60	86.677	1.00	0.0237	0.86	0.0213		
		86.0	0.0000	74.0	100.00	8.60	10.326	7.40	76.352	0.86	0.0121	0.74	0.0092		
		74.0	0.0000	63.0	100.00	7.40	13.417	6.30	62.935	0.74	0.0064	0.63	0.0028		
		63.0	0.0000	54.0	100.00	6.30	14.999	5.40	47.935	0.63	0.0021	0.54	0.0006		
		54.0	0.0000	46.0	100.00	5.40	16.094	4.60	31.841	0.54	0.0006	0.46	0.0001		
		46.0	0.0000	40.0	100.00	4.60	13.547	4.00	18.295	0.46	0.0001	0.40	0.0000		
		40.0	0.0000	34.0	100.00	4.00	11.255	3.40	7.0394	0.40	0.0000	0.34	0.0000		
		34.0	0.0000	29.0	100.00	3.40	3.2799	2.90	3.7595	0.34	0.0000	0.29	0.0000		
		29.0	0.0000	25.0	100.00	2.90	1.3355	2.50	2.4240	0.29	0.0000	0.25	0.0000		
		25.0	0.0000	22.0	100.00	2.50	0.8131	2.20	1.6109	0.25	0.0000	0.22	0.0000		
		22.0	0.0000	18.0	100.00	2.20	0.8995	1.80	0.7114	0.22	0.0000	0.18	0.0000		
		18.0	0.0000	16.0	100.00	1.80	0.3128	1.60	0.3985	0.18	0.0000	0.16	0.0000		
180	0.0000	160	100.00	16.0	0.0000	14.0	0.0000	14.0	0.0000	0.16	0.0000	0.14	0.0000		
160	0.0000	140	100.00	14.0	0.0000	12.0	0.0000	12.0	0.0000	0.14	0.0000	0.12	0.0000		
140	0.0000	120	100.00	12.0	0.0000	10.0	0.0000	10.0	0.0000	0.12	0.0000	0.10	0.0000		
120	0.0000	100	100.00	10.0	5.8201	10.0	94.180	1.20	0.0613	1.00	0.0450				



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FIGURE 7(A)

Spray-dried nanoparticulate budesonide

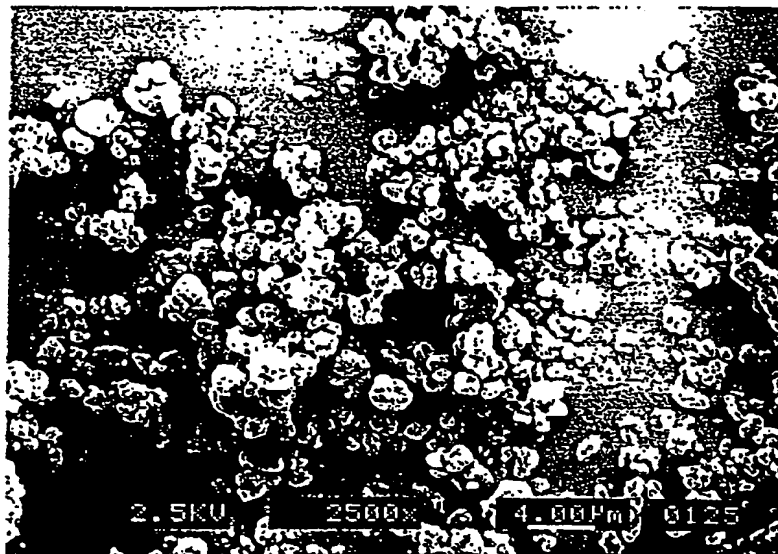


FIGURE 7(B)

Micronized budesonide



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FIGURE 8

HORIBA LA-910

Laser scattering particle size distribution analyzer

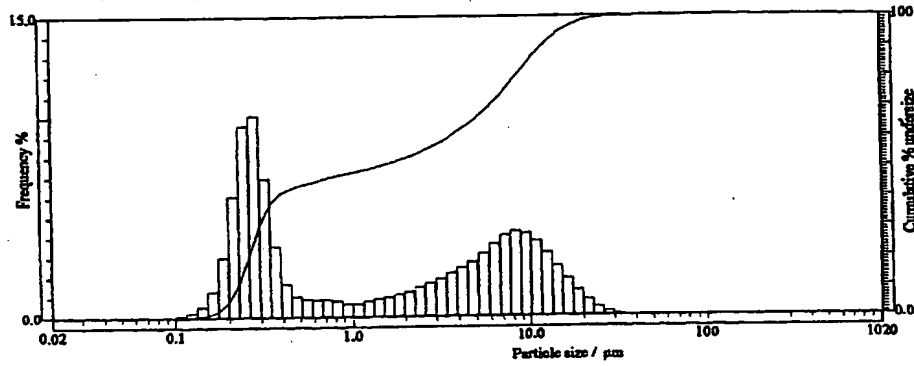
PARTICLE SIZE MEASUREMENT DATA

Freeze-dried Material : 5%Dextrose 4-3-98
Source : Reconst. water/3d fill
Lot Numbe : in water/1 min. sonication

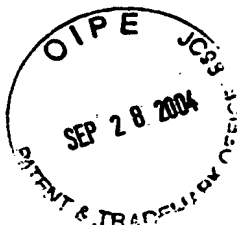
Data
Median : 1.355µm SP.Area: 114884cm²/cm³ S.D. : 5.324µm
Mode : 0.272µm Mean : 4.226µm
C.V. : 126.02%

Span : (D 10.0-D 90.0) / D50 = 8.564

Dia. on % (90.0%) :	11.622µm	% on Dia. (0.400µm) :	41.8%
Dia. on % (50.0%) :	1.355µm	% on Dia. (0.300µm) :	31.7%
Dia. on % (95.0%) :	14.996µm	% on Dia. (0.100µm) :	0.0%
Dia. on % (80.0%) :	8.384µm	% on Dia. (0.200µm) :	5.9%
Dia. on % (70.0%) :	5.949µm	% on Dia. (1.000µm) :	48.3%



Size(µm)Freq(%Und(%))			Size(µm)Freq(%Und(%))			Size(µm)Freq(%Und(%))		
1019.5	0.00	100.00	26.11	0.46	99.66	0.669	0.66	46.02
890.1	0.00	100.00	22.80	0.81	99.20	0.584	0.90	45.16
777.1	0.00	100.00	19.90	1.29	98.39	0.510	1.03	44.26
676.5	0.00	100.00	17.38	1.88	97.10	0.446	1.68	43.23
592.4	0.00	100.00	15.17	2.54	95.22	0.389	3.56	41.54
517.2	0.00	100.00	13.25	3.20	92.66	0.339	6.97	37.98
451.6	0.00	100.00	11.66	3.76	89.48	0.298	10.10	31.01
394.2	0.00	100.00	10.10	4.15	85.72	0.259	9.61	20.91
344.2	0.00	100.00	8.816	4.26	81.57	0.226	6.06	11.30
300.5	0.00	100.00	7.697	4.06	77.32	0.197	2.99	5.23
262.4	0.00	100.00	6.720	3.63	73.26	0.172	1.31	2.24
229.1	0.00	100.00	5.867	3.16	69.63	0.150	0.66	0.93
200.0	0.00	100.00	5.122	2.76	66.47	0.131	0.26	0.37
174.6	0.00	100.00	4.472	2.46	63.71	0.115	0.11	0.11
152.5	0.00	100.00	3.905	2.16	61.25	0.100	0.00	0.00
133.1	0.00	100.00	3.408	1.89	59.10	0.087	0.00	0.00
116.2	0.00	100.00	2.976	1.68	57.21	0.076	0.00	0.00
101.5	0.00	100.00	2.599	1.49	55.52	0.067	0.00	0.00
88.58	0.00	100.00	2.269	1.24	54.03	0.058	0.00	0.00
77.34	0.00	100.00	1.981	1.10	52.79	0.051	0.00	0.00
67.52	0.00	100.00	1.729	0.97	51.69	0.044	0.00	0.00
58.95	0.00	100.00	1.510	0.90	50.72	0.039	0.00	0.00
51.47	0.00	100.00	1.318	0.80	49.82	0.034	0.00	0.00
44.84	0.00	100.00	1.151	0.67	49.02	0.029	0.00	0.00
39.23	0.00	100.00	1.005	0.66	48.35	0.026	0.00	0.00
34.25	0.11	100.00	0.877	0.81	47.69	0.022	0.00	0.00
29.91	0.24	99.89	0.766	0.86	46.88			



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FIGURE 9

HORIBA LA-910

Laser scattering particle size distribution analyzer

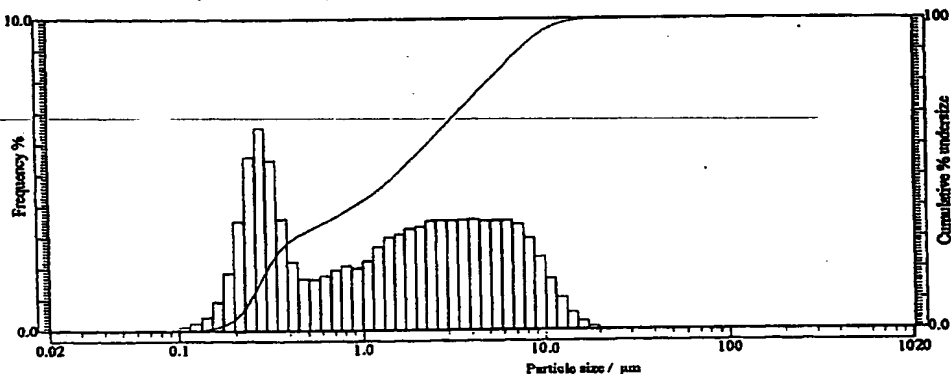
PARTICLE SIZE MEASUREMENT DATA

Material : reconst. 1%N9588, 5%Man
Source : in water
Lot Number : 1min sonication

Data
Median : 1.533 μ m SP.Area: 93485cm²/cm³ S.D. : 3.123 μ m
Mode : 0.276 μ m Mean : 2.767 μ m
C.V. : 112.86%

Span : (D 10.0-D 90.0) / D50 = 4.665

Dia. on % (90.0%) :	7.392 μ m	% on Dia. (0.400 μ m) :	28.8%
Dia. on % (50.0%) :	1.533 μ m	% on Dia. (0.300 μ m) :	19.8%
Dia. on % (95.0%) :	9.346 μ m	% on Dia. (0.100 μ m) :	0.0%
Dia. on % (80.0%) :	5.011 μ m	% on Dia. (0.200 μ m) :	4.0%
Dia. on % (70.0%) :	3.416 μ m	% on Dia. (1.000 μ m) :	41.7%

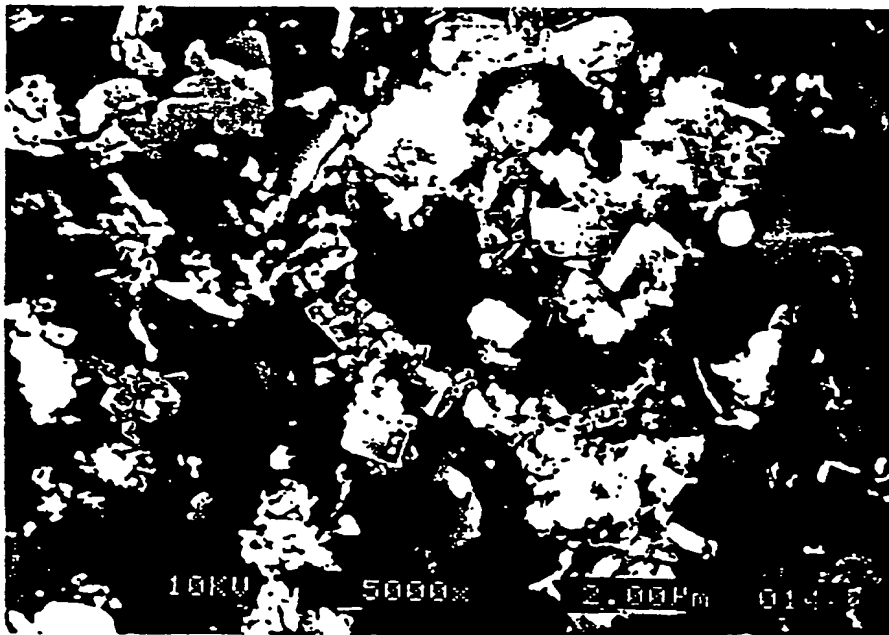


Size(μ m)	Freq(%)	Und(%)	Size(μ m)	Freq(%)	Und(%)	Size(μ m)	Freq(%)	Und(%)
1019.5	0.00	100.00	28.11	0.00	100.00	0.669	1.75	35.86
890.1	0.00	100.00	22.80	0.00	100.00	0.584	1.65	33.91
777.1	0.00	100.00	19.90	0.11	100.00	0.510	1.67	32.26
678.5	0.00	100.00	17.38	0.27	99.89	0.445	2.22	30.59
592.4	0.00	100.00	15.17	0.57	99.61	0.389	3.60	28.36
517.2	0.00	100.00	13.25	1.04	99.04	0.339	5.49	24.76
451.6	0.00	100.00	11.56	1.66	98.00	0.296	6.54	19.28
394.2	0.00	100.00	10.10	2.35	96.34	0.259	5.59	12.74
344.2	0.00	100.00	8.816	2.98	93.99	0.226	3.52	7.16
300.6	0.00	100.00	7.697	3.38	91.01	0.197	1.87	3.63
262.4	0.00	100.00	6.720	3.53	87.82	0.172	0.93	1.76
229.1	0.00	100.00	5.867	3.52	84.09	0.150	0.46	0.64
200.0	0.00	100.00	5.122	3.51	80.57	0.131	0.25	0.38
174.6	0.00	100.00	4.472	3.55	77.06	0.115	0.13	0.13
152.5	0.00	100.00	3.905	3.56	73.50	0.100	0.00	0.00
133.1	0.00	100.00	3.409	3.53	69.94	0.087	0.00	0.00
116.2	0.00	100.00	2.976	3.54	66.41	0.076	0.00	0.00
101.5	0.00	100.00	2.599	3.51	62.87	0.067	0.00	0.00
88.58	0.00	100.00	2.269	3.33	59.36	0.058	0.00	0.00
77.34	0.00	100.00	1.981	3.27	56.03	0.051	0.00	0.00
67.52	0.00	100.00	1.729	3.10	52.76	0.044	0.00	0.00
58.95	0.00	100.00	1.510	3.00	49.66	0.039	0.00	0.00
51.47	0.00	100.00	1.318	2.70	46.86	0.034	0.00	0.00
44.94	0.00	100.00	1.151	2.24	43.96	0.029	0.00	0.00
39.23	0.00	100.00	1.005	2.02	41.73	0.026	0.00	0.00
34.25	0.00	100.00	0.877	2.09	39.71	0.022	0.00	0.00
29.91	0.00	100.00	0.766	1.95	37.62			



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FIGURE 10



**Micrograph of
Milled TA (3.6%) with Span 85 (0.5%)**